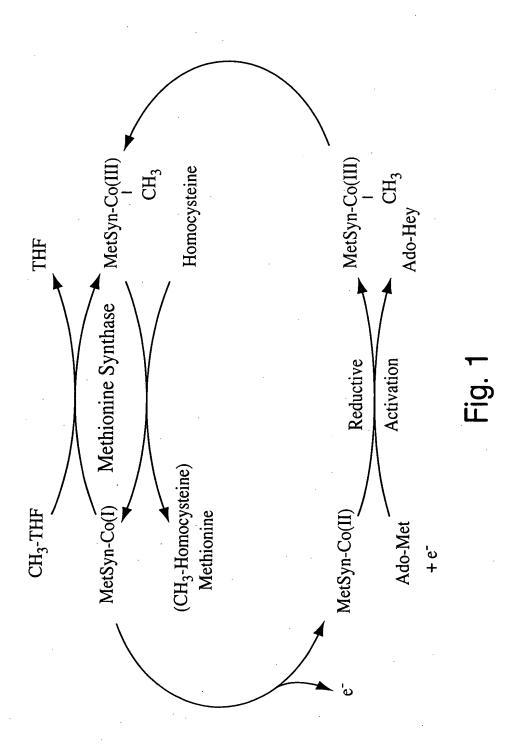


REPLACEMENT SHEET
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REPLACEMENT SHEET

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CAAGGTTGGTGGAAGTCGCGTTGTGCAGGTTCGTGCCCGGCTGGCGGCGTGGTTTCACTGTTACATGCCTTGAAGTG

ATGAGGAGGTTTCTGTTACTATATGCTACAGCAGGGACAGGCAAAAGGCCATCGCAGAAAAATGTGT

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Fig. 3-

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Fig. 3-4

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HsMTRR		
CeMTRR		
HsCPR	MGDSHVDTSSTVSEAVAEEVSLFSMTDMILFSLIVGLLTYWFLFRKKKEE FMN	5(
HsMTRR	MPDFI.T.VATOOCOAVATAREMOR	24
CeMTRR	MTDFLIAFGSQTGQAETIAKSLKE	24
HsCPR	VPEFTKIQTLTSSVRESSFVEKMKKTGRNIIVFYGSQTGTAEEFANRLSK	100
	FMN	
HsMTRR	QAVVHGFSADLHCISESDK-YDLKTETAPLVVVVSTTGTGDPP	66
CeMTRR	KAELIGLTPRLHALDENEKKFNLNEEKLC-AIVVSSTGDGDAP	66
HsCPR	DAHRYGMRGMSADPEEYDLADLSSLPEIDNALVVFCMATYGEGDPT	146
	* *	11(
	FMN	
HsMTRR	DTARKFVKEIQNQTLPVDFFAHLRYGLLGLGDSEYTYFCNGGKIIDKRLQ	116
CeMTRR	DNCARFVRRINRNSLENEYLKNLDYVLLGLGDSNYSSYQTIPRKIDKQLT	116
HsCPR	DNAQDFYDWLQETDVDLSGVKFAVFGLGNKTYEHFNAMGKYVDKRLE	193
	* *	
HsMTRR	ELGARHFYDTGHADDCVGLELVVEPWIAGLWPALRKHFRSSRGQEEISGA	166
CeMTRR	ALGANRLFDRAEADDQVGLELEVEPWIEKFFATLASRFDISADKMN	162
HsCPR	QLGAQRIFELGLGDDDGNLEEDFITWREQFWPAVCEHFGV	233
·	*** ** ** * *	. 433
HsMTRR	LPVASPASLRTDLVKSELLHIESQVELLRFDDSGRKDSEVLKQNAVNS	214
CeMTRR	-AITESSNLKLNQVKTEEEKKALLQKRIEDEESDDEGRGRVIGID-	206
HsCPR	EATGEESSIRQYELVVHTDIDAAKVYMGEMGRLKSYEN	271
	• • • • • • • • • • • • • • • • • • • •	
HsMTRR	NQSNVVIEDFESSLTRSVPPLS-QASLNIPGLPPEYLQVHLQESLGQ	260
CeMTRR	MLIPEHYDYPEISLLKGSQTLSNDENLRVPIAPQPFIVSSVSNRKLP	253
HsCPR	QKPPFDAKNPFLAAVTTNRKLN	293
	• • • •	233
HsMTRR	EESQVSVTSADPVFQVPISKAVQLTTNDAIKTTLLVELDIS	301
CeMTRR	EDTKLEWQNLCKMPGVVTKPFEVLVVSAEFVTDPFSKKIKTKRMITVDFG	303
HsCPR	QGTERHLMHLELD	306
HsMTRR	NTDFSYQPGDAFSVICPNSDSEVQSLLQR-LQLEDKREHCVLLKIKAD	348
CeMTRR	DHAAELQYEPGDAIYFCVPNPALEVNFILKRCGVLDIADQQCEL-SINPK	352
HsCPR	ISDSKIRYESGDHVAVYPANDSALVNQLGKILGADLDVVMSLNNL	351
	. *. **	771
HsMTRR	TKKKGATLPQHIPAGCSLQFIFTWCLEIRAIPKKAFLRALVDYTSDSAEK	398
CeMTRR	TEKINAQIPGHVHKITTLRHMFTTCLDIRRAPGRPLIRVLAESTSDPNEK	402
HsCPR	DEESNKKHPFPCPTSYRTALTYYLDITNPPRTNVLYELAQYASEPSEQ	399
	+ + + + + + + + + + + + + + + + + + +	555

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HsMTRR	RRLQELCSKQGAADYSRFVRDACACLLDLLLAFPSCQPPLSLLLEHLP	446
CeMTRR	RRLLELCSAOGMKDFTDFVRTPGLSLADMLFAFPNVKPPVDRLIELLP	450
• •	ELLRKMASSSGEGKELYLSWVVEARRHILAILQDCPSLRPPIDHLCELLP	449
HsCPR		777
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	FAD FAD FAD	
HsMTRR	KLQPRPYSCASSSLFHPGKLHFVFNIVEFLSTATTEVLRKGVCTGWLALL	496
CeMTRR	RLIPRPYSMSSYENRKARLIYSEMEFPATDGRRHSRKGLATDWLNSL	497
HsCPR	RLQARYYSIASSSKVHPNSVHICAVVVEYETKAGRINKGVATNWL	494
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TI-MMDD	TINCIT ODNITUR CUEDCOVAL ADVICTODD TONICEULD DODGED	539
HsMTRR	VASVLQPNIHASHEDSGKALAPKISISPRTTNSFHLPDDPSIP	
CeMTRR	RIGDKVQVLGKEPARFRLPPLGMTKNSAGKLP	529
HsCPR	RAKEPVGENGGRALVPMFVRKSQFRLPFKATTP	527
	*	
	NADPH NADPH	
HsMTRR	IIMVGPGTGIAPFIGFLQHREKLQEQHPDGNFGAMW-LFFGCRHKDRDYL	588
CeMTRR	LLMVGPGTGVSVFLSFLHFLRKLKQDSPSDFVDVPRVLFFGCRDSSVDAI	579
HsCPR	VIMVGPGTGVAPFIGFIQERAWLRQQGKEVGETLLYYGCRRSDEDYL	574
	. * * * * * * * * * * * * * * * * * * *	4
	NADPH NADPH	
HsMTRR	FRKELRHFLKHGILTHLKVSFSRDAPVGEEEAPAKYVQDNIQLHGQQVAR	638
CeMTRR	YMSELEMFVSEGILTDLIICESEQKGERVQDGLRKYLDKVLP	621
HsCPR	YREELAQFHRDGALTQLNVAFSREQSHKVYVQHLLKQDREHLWK	618
	. ** * * ** * . * *	
•	NADPH	
HsMTRR	ILLQE-NGHIYVCGDAKNMAKDVHDALVQIISKEVGVEKLEAMKTLATLK	687
CeMTRR	FLTASTESKIFICGDAKGMSKDVWQCFSDIVASDQGIPDLEAKKKLMDLK	671
HsCPR	LIEGGAHIYVCGDARNMARDVQNTFYDIVAELGAMEHAQAVDYIKKLM	666
•		
	NADPH/FAD	
HsMTRR	EEKRYLODIWS 698	
CeMTRR	KSDQYIEDVWG 682	
HsCPR	TKGRYSLDVWS 677	
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Fig. 4-2

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9/14. **HsMTRR CeMTRR** MGDSHVDTSSTVSEAVAEEVSLFSMTDMILFSLIVGLLTYWFLFRKKKEE HsCPR 50 FMN -----MRRFLLLYATQQGQAKAIAEEMCE **HsMTRR** 24 -----MTDFLIAFGSQTGQAETIAKSLKE CeMTRR 24 HsCPR VPEFTKIQTLTSSVRESSFVEKMKKTGRNIIVFYGSQTGTAEEFANRLSK 100 FMN QAVVHGFSADLHCISESDK-YDLKT-----ETAPLVVVVSTTGTGDPP **HsMTRR** 66 **CeMTRR** KAELIGLTPRLHALDENEKKFNLNE-----EKLC-AIVVSSTGDGDAP 66 **HsCPR** DAHRYG----MRGMSADPEEYDLADLSSLPEIDNALVVFCMATYGEGDPT 146 **HsMTRR** DTARKFVKEIQNQTLPVDFFAHLRYGLLGLGDSEYTYFCNGGKIIDKRLO 116 **CeMTRR** DNCARFVRRINRNSLENEYLKNLDYVLLGLGDSNYSSYQTIPRKIDKQLT 116 **HsCPR** DNAQDFYDWLQETDVD---LSGVKFAVFGLGNKTYEHFNAMGKYVDKRLE 193 **HsMTRR** ELGARHFYDTGHADDCVGLELVVEPWIAGLWPALRKHFRSSRGQEEISGA 166 CeMTRR ALGANRLFDRAEADDOVGLELEVEPWIEKFFATLASRFDISADKMN----162 HsCPR QLGAQRIFELGLGDDDGNLEEDFITWREQFWPAVCEHF-----GV 233 HsMTRR LPVASPASLRTDLVKSELLHIESQVELL--RFDDSGRKDSEVLKQNAVNS 214 CeMTRR -AITESSNLKLNQVKTE----EEKKALLQKRIEDEESDDEGRGRVIGID-206 EATGEESSIRQYEL-------VVHTDIDAAKVYMGEMGRLKSYEN HsCPR 271 NQSNVVIEDF---ESSLTRSVPPLS-QASLNIPGLPPEYLQVHLQESLGQ HSMTRR 260 ---MLIPEHYDYPEISLLKGSQTLSNDENLRVPIAPQPFIVSSVSNRKLP CeMTRR 253 **HsCPR** OKP-----PFD-------AKNPFLAAVTTNRKLN 293 **HsMTRR** EESQVS------VTSADPVFQVPISKAVOLTT--NDAIKTTLLVELDIS 301 CeMTRR EDTKLEWQNLCKMPGVVTKPFEVLVVSAEFVTDPFSKKIKTKRMITVDFG 303 OGTE-----RHLMHLELD HsCPR 306 HSMTRR N--TDFSYQPGDAFSVICPNSDSEVQSLLQR-LQLEDKREHCVLLKIKAD 348 CeMTRR DHAAELQYEPGDAIYFCVPNPALEVNFILKRCGVLDIADOOCEL-SINPK 352 **HsCPR** ISDSKIRYESGDHVAVYPANDSALVNOLGK---ILGADLD--VVMSLNNL **HSMTRR** TKKKGATLPQHIPAGCSLQFIFTWCLEIRAIPKKAFLRALVDYTSDSAEK 398 CeMTRR TEKINAQIPGHVHKITTLRHMFTTCLDIRRAPGRPLIRVLAESTSDPNEK 402 HsCPR DEESNKKHP--FPCPTSYRTALTYYLDITNPPRTNVLYELAQYASEPSEO 399

Fig. 4-3

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HsMTRR	RRLQELCSKQGAADYSRFVRDACACLLDLLLAFPSCQPPLSLLLEHLP	446
CeMTRR	RRLLELCSAQGMKDFTDFVRTPGLSLADMLFAFPNVKPPVDRLIELLP	450
• •	——————————————————————————————————————	
HsCPR	ELLRKMASSSGEGKELYLSWVVEARRHILAILQDCPSLRPPIDHLCELLP	449
	* . * .*** * **. * **.	
	FAD FAD FAD	
HsMTRR	KLQPRPYSCASSSLFHPGKLHFVFNIVEFLSTATTEVLRKGVCTGWLALL	496
CeMTRR	RLIPRPYSMSS YENRKARLIYSEMEFPATDGRRHSRKGLATDWLNSL	
		497
HsCPR	RLQARYYSIASSSKVHPNSVHICAVVVEYETKAGRINKGVATNWL	494
	.* * ** *	
HsMTRR	VASVLQPNIHASHEDSGKALAPKISISPRTTNSFHLPDDPSIP	539
CeMTRR	RIGDKVQVLGKEPARFRLPPLGMTKNSAGKLP	529
* *		
HsCPR	RAKEPVGENGGRALVPMFVRKSQFRLPFKATTP	527
	*	
	NADPH NADPH	
HsMTRR	IIMVGPGTGIAPFIGFLQHREKLQEQHPDGNFGAMW-LFFGCRHKDRDYL	588
CeMTRR	LLMVGPGTGVSVFLSFLHFLRKLKQDSPSDFVDVPRVLFFGCRDSSVDAI	579
HsCPR	VIMVGPGTGVAPFIGFIQERAWLRQQGKEVGETLLYYGCRRSDEDYL	574
naci k		3/4
	•	
	NADPH NADPH	
HsMTRR	FRKELRHFLKHGILTHLKVSFSRDAPVGEEEAPAKYVQDNIQLHGQQVAR	638
CeMTRR	YMSELEMFVSEGILTDLIICESEQKGERVQDGLRKYLDKVLP	621
HsCPR	YREELAQFHRDGALTQLNVAFSREQSHKVYVQHLLKQDREHLWK	618
	** * * * * * * * * * *	010
	NADPH	
tt = MMD D		
HsMTRR	ILLQE-NGHIYVCGDAKNMAKDVHDALVQIISKEVGVEKLEAMKTLATLK	687
CeMTRR	FLTASTESKIFICGDAKGMSKDVWQCFSDIVASDQGIPDLEAKKKLMDLK	671
HsCPR	LIEGGAHIYVCGDARNMARDVQNTFYDIVAELGAMEHAQAVDYIKKLM	666
	NADPH/FAD	
HsMTRR	EEKRYLQDIWS 698	
CeMTRR	KSDQYIEDVWG 682	
	•••	
HsCPR	TKGRYSLDVWS 677	
	* * *	

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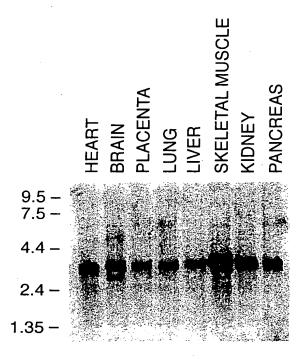


Fig. 5A



Fig. 5B

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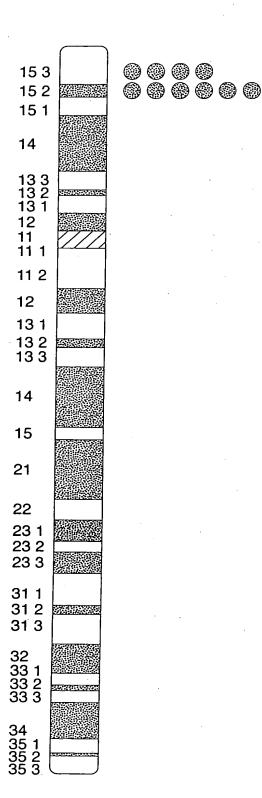


Fig. 6

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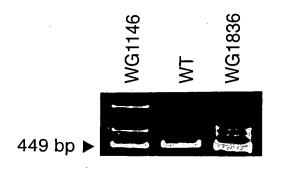


Fig. 7A

WG1836

336 bp ▶

Fig. 7B

Position	Sequence	Accession#	Protein		Organism	
572	GAMWLFFGCRHKDRDYLF	(AF025794)	MTRR		(H sapiens)	
558	GETL L YYGCRRSDEDYLY	(A60557)	CPR		(H sapiens)	
559	GETLLYYGCRRAAEDYLY	(D00101)	CPR		(O cuniculus)	
560	GESILYFGCRKRSEDYIY	(X93090)	CPR		(D melanogaster)	
572	GPALLFFGCRNRQMDF1Y	(P37116)	CPR		(V radiata)	
573	GPTV L FFGCRKSDEDFLY	(Z26938)	CPR		(A niger)	
1281	CPMV L VFGCRQSKIDHIY	(D16408)	NOS	I	(H sapiens)	
1009	GRMTLVFGCRRPDEDHIY	(U05810)	NOS	ΙΙ	(H sapiens)	
1040	TPMTLVFGCRCSQLDHLY	(L26914)	NOS	III	(H sapiens)	
380	GRMTLVFGCRHPEEDHLY	(U85094)	NOS		(O cuniculus)	
1005	GDMILLFGCRHPDMDHIY	(U46504)	NOS		(G gallus)	
481	GKNWLFFGNPHFTEDFLY	(M23008)	SR		(E coli)	
915	GEVFLYLGSRHKREEYLY	(L26503)	SR		(S cerevisiae)	
407	GRNWLIFGNRHFHRDFLY	(Z23169)	SR		(T roseopersicina)	
261	GLAWLFLGVANVDSLLYD	(X99419)	FNR		(P sativum)	
251	GLAWLFLGVPTSSSLLYK	(P00455)	FNR		(S oleracea)	

Fig. 7C

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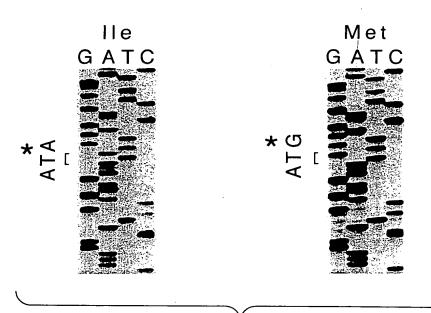


Fig. 8A

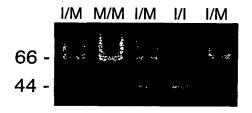


Fig. 8B